P. C.	T RAI	ATRADEMARK OF			1/59					
	>	FIG. 1AG	FIG. 1AH	FIG. 1AI	FIG. 1AJ	FIG. 1AK	FIG. 1AL	FIG. 1AM	FIG. 1AN	
65	>	FIG. 1Y	FIG. 1Z	FIG. 1AA	FIG. 1AB	FIG. 1AC	FIG. 1AD	FIG. 1AE	FIG. 1AF	
se Page 1 of 59		ď	1R		L	n	Λ.	1W	1X	
Polycystic Kidney Disease		FIG. 1Q	FIG. 1	FIG. 1S	FIG. 1T	FIG. 1U	FIG. 1V	FIG. 1	FIG. 1	
Pol			 -			· ·				
		11	11	1K	1T	1 M	1N	10	11	
		FIG.	FIG. 1J	FIG. 1K	FIG. 1L	FIG. 1M	FIG. 1N	FIG. 10	FIG.	
ī	L									<u>-</u>
		1A	1B	1C	1D	1E	1F	1G	1H	
	÷	FIG. 1A	FIG. 1B	FIG. 1C	FIG. 1D	FIG. 1E	FIG. 1F	FIG. 1G	FIG. 1H	

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Docket No.: GZ 2061.00 U.S. Serial No.: 09/830,506 Applicant: Oxana Ibraghimov-Beskrovnaya, et al. Title: Compositions and Methods for Treating Polycystic Kidney Disease

FIG. 1



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Polycystic Kidney Disease
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								1
	09	120	170	218	266	314	362	410
Polycystic Kidney Disease Page 2 of 59	GETCECEGC CECAGCCCCA TCCAGCCCGC GCCCGCCATG CCGTCCGCGG	AGCTGCGGCC TCCGCGCGCGC GCGGCCTG GGGACGGCGG GGCCATGCGC	AACG ATG CCG CCC GCC CCC CGC CTG GCG CTG GCC Met Pro Pro Aid Aid Pro Aid Arg Leu Aid Leu Aid 5	CTG GGC CTG TGG CTC GGG GCG GGG GGC CCC GGG CGC 2 Leu Gly Leu Trp Leu Gly Ala Leu Ala Gly Gly Pro Gly Arg 15	GGG CCC TGC GAG CCC CCC TGC CTC TGC GGC CCA GCG CCC GGC 2 Gly Pro Cys Glu Pro Pro Cys Leu Cys Gly Pro Alg Pro Gly 35	TGC CGC GTC AAC TGC TCG GGC CGC CTG CGG ACG CTC GGT 3 Cys Arg Val Asn Cys Ser Gly Arg Gly Leu Arg Thr Leu Gly 50 55 60	CGC ATC CCC GCG GAC GCC ACA GCG CTA GAC GTC TCC CAC Arg ile Pro Ala Asp Ala Thr Ala Leu Asp Val Ser His 65 75	cGG GCG CTG GAC GTT GGG CTC CTG GCG AAC CTC TCG GCG Arg Aid Leu Asp Vai Gly Leu Leu Aid Asn Leu Ser Aid 80
	GCAG	GCCT	၁၁၁၅	1 KI9	TGC G Cys G 30	200	GCG CTG Ala Leu) 10 C
·	GCTCAGCAGC	910000000	ССССТССССТ	CTG G	66C T 617 C	GCC GCC Ala Ala 45	Pro A	AAC CTG CTC Asn Leu Leu

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APR 0 1 2003	JC48			•		
PADEMARK OFF			3/59			1B
			·	·		FIG.
458	206	554	602	650	869	746
GAA	666 919	6CG A I a 140	GCT	CTG Fen	AGC	CTG Leu
GAA	AGT Ser	166 Trp	TGT Cys 155	TTG	AAC Asn	660 617
TTA Leu	CTG Leu	CGA Arg	ACG Thr	CCC Pro 170	GAC ASP	GAA Glu
ACG Thr 105	AAC Asn	cce Pro	D I V	ATC I le	CCT Pro 185	CAC
Ser	ATA Ile 120	CTG Leu	GCA	660 61 y	CTC	GCC A I d 200
ATT IIe	GAA	TGG Trp 135	GAG G1u	CTT	TGC Cys	GCT
AAG Lys	AGT Ser	000 A I a	CCC Pro 150	CTG	BIA	TCA
AAC	TTA	CTG	CAG GIn	CCT Pro 165	GTC	Phe
AAC ASN 100	AAT Asn	660 61 y	616 Val	CAG GIn	TAT Tyr 180	TCC Ser
AGC Ser	TTT Phe 115	TGT Cys	6T6 Val	660 61y	GAG G I u	GTG Val 195
ATA Ile	TTA	GAC Asp 130	CGG Arg	GCT	GAG GIU	GCA A I d
GAT Asp	AAT Asn	TGT Cys	GTG Val 145	CTG	66T 61y	GCA A I d
CTG	GCT	GAG	CAG	TCC Ser 160	16T Cys	GTG Val
GAG Glu 95	TTT	TTT	CAG GIn	66 617	660 61 y 175	ACC Thr
GCA	ATA 116 110	CCG Pro	GAG	CCT Pro	AGT Ser	66C 61√ 190
CTG	66A 61y	AAC Asn 125	GAG	666 61 y	GAC Asp	TCA Ser

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		•			
	794	. 842	890	938	986
	66C 61 y 220	CAG GIN	cc6 Pro	GTC Val	CTG
Applicant: Oxana Braghimov-Beskrovnaya, et al. Title: Compositions and Methods for Treating Polycystic Kidney Disease Page 4 of 59	CAG	6CC A10 235	000 P. C	CAC His	CCT Pro
a, et al. ing of 59	ح ب	B A	660 61y 250	CAG	GGA GIY
skrovnay for Treati Page 4 o	ACC	666 61 y	TCC Ser	CTC Leu 265	CAC His
imov-Be Methods	TCC Ser	TGT Cys	16c Cys	CTC	CCC Pro 280
Applicant: Oxana Braghime Title: Compositions and Met Polycystic Kidney Disease	TTC Phe 215	CTG	CTC	ACC Thr	666 61 y
cant: Oxa Composit stic Kidn	sko Cys	TGC Cys 230	Ser	000 Pro	GTG Val
Applic Title: (Polycy	TTC Phe	166 1rp	CTG Leu 245	66c	CTG CTG
	B I A	660 61 y	16c Cys	AGG Arg 260	ACC Thr
	AGC	CAG GIN	BIA	161 Cys	GCC A10 275
	TGC Cys 210	GAG Glu	TTT Phe	ACC Thr	666 61y
	000 V I a	TCG Ser 225	Ser	CCC Pro	CCA Pro
	GAG Glu	CTC	GCC A10 240	BIA	TCC
	CCA	B I A	AGT Se r	CCT Pro 255	GCC
	CAG	GCA A I a	TCC	CCT Pro	CCT Pro 270
	CTT Leu 205	CTC	CCC Pro	CCA	TTC Phe
	1				

FIG. 1C

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GTG GAT GCC Val Asp Ald 315

TCC GCC GAG C

GGA GAC GGC 1 Gly Asp Gly 5

Phe

Asp

TGG Trp 305

ACA CGC Thr Arg

OCC A

ACT Thr

GAC TTC

1034

CIC ren

ပ္ပ Pro

ပ္ပ AIG

CAC ATC

GCT Ald 295

Ile

His

TTC (

CTA GCA GCC :

GGC CAG

101 Ser

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Gin

GIY

A10 285

A I a 290

CCT GTC Pro Val 300

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APR 0 1 2003	ر د د د د د د د د د د د د د د د د د د د		5/59			1D
						FIG.
1130	1178	1226	1274	1322	1370	1418
CAC His	ACA Thr	cce Pro	CGC Arg 380	GAG G I u	ATC Ile	BIA
rat Tyr	666 61 y	16C Cys	AAC	66C 61 y 395	GAG G I u	B I A
CGC Arg 330	CTG Leu	6T6 Val	CAG	CTG	ACG Thr 410	AAG Lys
666 61y	CTG Leu 345	CTC	ATC I le	GCC	GAC Asp	GAG G1u 425
Pro (D I V	GAG G1u 360	AGC ATC Ser Ile	GTG Val	TCG	6T6 Vai
CTG	Ser	CTG	CTC Leu 375	ATC Ile	CCC Pro	616 Va i
GTG Val	66C 61y	B I A	GAC Asp	AGC Ser 390	16C Cys	CTG Leu
TAT Tyr 325	D I V	D I V	CTC	r AC Ty r	CTC Leu 405	CGC
SGC Arg	666 61 y 340	Pro	AGC Se r	DIA	CCG Pro	TAC Tyr 420
CAT (His A	CTG Leu	GCA A10 355	GAG Glu	B A	CAC His	760 Cys
TCG Ser	B I A	BLA Ald	GAC Asp 370	GAG Glu	GTG Val	CAC His
SCC A I a	CTG	GAA Glu	AGT Ser	CTG Leu 385	GCG	666 61 y
GCT Ald 320	GTG Val	GTG Val	CAG GIn	660 61 y	CGA Arg 400	AAC
CCG Pro	GCC A I a 335	CAG	616 Val	TCA	BIA Ala	GGC G1 y 415
666 61 y	ACG Thr	6T6 Val 350	TCG Ser	66T 61 y	CCG Pro	CCT Pro
GCT	GTG	GAC ASP	TCC Ser 365	661 61 y	GAG Glu	TTC Phe

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	ADEMARK OHIER			6/59			1E
.					·		FIG.
	1466	1514	1562	1610	1658	1706	1754
 	CTG	GTC Val 460	GTG Val	16C Cys	TGC Cys	000 910	GAT ASP 540
I. I I	B I A	CGG	666 61 y 475	AGC	CAC His	TCA Ser	CAG GIn
rig 159 1	BIA	TCC Se r	CAG GIn	GAG Glu 490	GAG Glu	76C Cys	GTG Vai
or Treating Page 6 of 59	666 61 y	GTC Val	6T6 Val	CTG	GCC A1a 505	CTG	CCA Pro
Title: Compositions and Methods for Treating Polycystic Kidney Disease Page 6 of 5	6CC A 1 a	CTG	ACT	AGC Ser	ACA Thr	GAC ASP 520	66c
ns and My Disease	166 Trp	TTC Phe 455	TCG Ser	TTC Phe	BIA	ACC Thr	GGA GLY 535
ompositio tic Kidney	GCC TGG Ald Trp	CGC Arg	TTC Phe 470	DIA	CCA Pro	AAC Asn	CCC Pro
Title: Co	CAG	CAG	66C 61y	GAG G1u 485	CAC His	TGT Cys	CAG GIn
	TGT Cys	GTG Val	ATC Ile	66C 61y	CCA Pro 500	166 1rp	CTG
j	CAG Gin 435	B A	766 Trp	CAG GIn	GAG Glu	666 61 y 515	GAG
; 	GAG Glu	CCC Pro 450	616 Val	CCG Pro	666 61y	ACC	TGC Cys 530
j 1	CAG GIn	AGT	GAC ASP 465	GCG A I a	CCC Pro	CCC Pro	GTC Val
	GCG A I a	GAC ASP	TGC CTA Cys Leu	CCA Pro 480	CTG	666 61 y	TAC
	CAG GIN	GTG Val	TGC Cys	660 61y	TGG Trp 495	CTC	AGC Se r
1 1	CTG Leu 430	ATG Met	AGG Arg	6T6 Val	AAC	CGG Arg 510	CAC His
 	166 Trp	GCA A10 445	ACC Thr	GAG G1u	CAG GIn	GTC Val	CCG Pro 525

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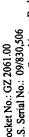
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						FIG
1802	1850	1898	1946	1994	2042	2090
CCC Pro	000 P 70	TTC	CAG	GAG G1u 620	CTG Leu	ATC Ile
GGA CCC G1y Pro 555	GAG GI u	B I A	CCC GCC Pro Ala	CCG Pro	CAG G1n 635	GCC AAC A Ala Asn 1 650
SAG 3 I n	CAC His 570	GAA Glu	CCC Pro	ACC (Thr	ACC	650 650
CTG	CCG Pro	CGT Arg 585	SGG Arg	666 61 y	AGG Arg	66A 61 y
GAC	B I A	AGC	CGG Arg 600	GCA	AAC Asn	TGC CCT (Cys Pro
GGG GAC CTG (Gly Asp Leu (TCA Se r	CTG	CTC CGG (Leu Arg / 600	ACA Thr 615	GAC AAC AGG Asp Asn Arg	16C Cys
CCC AGT (Pro Ser (550	GGC CTC 619 565	SGT Arg	GAG G I u	AGC) CG) TO 530	GGA CGC TGG Gly Arg Trp 645
CCC Pro	660 61 y 565	CTG (CAG GIn	CTC	AGG TCC (Arg Ser F	CGC Arg 645
B A	GAC Asp	660 61 y 580	ACC Thr	CTC	AGG Arg	66A 61 y
66A 61y	CAG	000 P 70	666 61 y 595	CGG Arg	AGC	666 61 <i>y</i>
GTG Val	CAG	TTC Phe	TTT Phe	GTG TAC Val Tyr 610	GAG AGC /	CCA Pro
CTC Leu 545	BCA Ald	GTA TTC CCG Val Phe Pro	GAA Glu	GTG Val	CCT Pro 625	ATG CCA GGG Met Pro Gly
CTC	CTG Leu 560	ATG Met	D I V	CAG	GAG G1u	TGC Cys 640
AAC Asn	CCT Pro	GTC Val 575	ACG Thr	CTG	AGC	BIA AIG
GAG G I u	ACG Thr	GAG G1u	ACC Thr 590	CGG Arg	66C 61y	000 P.7
B I A	CTG	6T6 V01	CTC	CTG Leu 605	AAC	BIA AIG

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MARKOT			8/59			7
						FIG. 1G
2138	2186	2234	2282	2330	2378	2426
66C 61 y	ACA Arg	GTC Val 700	66C 61 y	GCT	TCG Ser	TGC Cys
AAT Asn	76G 7rp	TCG Ser	GTT Val 715	CCG Pro	D I V	B I A
D I d	CTA	TAC Tyr	CTC	TCG Ser 730	AAC Asn	766 7 rp
GCC TGC (Alg. Cys / 665	B I B	CAG	GAC	CAC TGC His Cys	TCC GCC Ser Ald 745	GGC ACT TGG GCC T Gly Thr Trp Ala C 760
OCC A I d	TAT Tyr 680	B I A	GGT G·I y	CAC His	TCC Ser	660 61y 760
CAG GIn	CCC Pro	CCC P r o 695	CCT Pro	CTG	ne i	GAG G I u
000 Pro	BCC AIG	P ro	CTC Leu 710	CTC	CCG TAC Pro Tyr.	CTG
CAC His	666 61 y	666 61 y	ATG Met	GCC A10 725	CCG Pro	CAG GIn
TGC Cys 660	CCC Pro	000 P d	CTC	660 61 y	900 A10 740	B A B
TCC Se r	CTA Leu 675	ccc Pro	GTC Val	CCT Pro	CGG Arg	CCA Pro 755
B A	666 61 y	6TT Val 690	GAT Asp	66c	CCC Pro	TTG Leu
GAC ASP	CCA Pro	TCC	CAG GIN 705	GCT	GGT 61y	CAC His
CTG	666 61 y	TTC Phe	660 61y	GAC ASP 720	CCT Pro	CCC CAC Pro His
CCG Pro 655	TCA Ser	CTC	CAC	CAC	CAC His 735	CTG
TTG Leu	ACG Thr 670	TTC Phe	CTC	CAG GIn	660 61y	TGG Trp 750
TGC	16c Cys	GAG G1u 685	ACC Thr	TTG Leu	CCC Pro	TCA



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RADEMAR	OFF STATE OF THE S			9/59			1H
							FIG.
·	2474	2522	2570	2618	2666	2714	2762
	6T6 Val 780	TAT Tyr	TCC Ser	TAC Tyr	BIA Ala	CGC Arg 860	B I A
]. 	ACC	CGC Arg 795	CTC	ATC Ile	TCA	GCT	TGC CCT Cys Pro 875
tal.	CTC	666 61 y	AAC Asn 810	GTC Val	66C 61 y	ACG Thr	16C Cys
krovnaya, et a or Treating Page 9 of 59	CAG	CCT Pro	CAC	CGG Arg 825	AAC Asn	D V	GTC Val
6 nov-Beskr ethods for P	GAA	CTG	AGG Arg	CTG	ACC Thr 840	ACG Thr	AAT Asn
Docket No.: GZ 2061.00 U.S. Serial No.: 09/830,506 Applicant: Oxana Braghimov-Beskrovnaya, et al. Title: Compositions and Methods for Treating Polycystic Kidney Disease Polycystic Kidney Disease	ACG Thr 775	CGG Arg	TCC Ser	666 61y	000 P 70	6CC A l a 855	GAG G I u
No.: GZ 2 ial No.: C nt. Oxanz ompositio	B A	CTG Leu 790	616 Val	GCT	616 Val	AAC Asn	TTT Phe 870
Docket N U.S. Ser Applicar Title: Cc Polycyst	GCA A I a	66A 61y	660 61 y 805	6T6 Va i	TAC Tyr	B A	CGC Arg
	CTT	CCT Pro	AAT Asn	CCA Pro 820	CTC	661 61 y	B A
	CTG	AAC	66c	TCC	CGC Arg 835	TCT Ser	Ser Ald
i 	CGG Arg 770	occ Pro	616 Val	GTC Val	66C 61y	GAC ASP 850	GTC Vai
[!	CTG Leu	AGG Arg 785	GAG Glu	GTG Vai	GAC Asp	6T6 Val	AGT Ser 865
	GCC CTG Ald Leu	TTG	6CA A10 800	GAC Asp	CGC GAC Arg Asp	CAG	660 61 y
ĺ	TGT Cys	66C 61y	CGG Arg	TTT Phe 815	CCC Pro	CTC	666 61 y
i	B I A	CTG	GTC Val	AGC Ser	GCC A I a 830	GTG Val	CCT Pro
	CCT Pro 765	CTG Leu	GAG Glu	16C Cys	CCT	TTG Leu 845	766 Trp



3098 3002 3050 2906 2954 2858 2810 AGC Se r GTG AAG Lys CTG AGC Ser 940 GTG Val Leu Va ACC Thr CAG Gun GAC Asp CCC Pro 955 Asp ე ე ე CAC His AGC Ser GAT TAT Tyr AAC ASN 970 ASN 890 GAG Glu CIC Leu ACG Thr AGC Se r AAC Applicant: Oxana Ibraghimov-Beskrovnaya, et al. Page 10 of 59 AIG ATC Ile ATT I | e 985 ASN ည္ဟ TAC Tyr AAC Fitle: Compositions and Methods for Treating 666 61 y 905 ACC Thr GTC ACC GAG G1u Arg AGG Arg GCC Ald 920 ည္ဟ Thr Val GAG GTu Docket No.: GZ 2061.00 U.S. Serial No.: 09/830,506 AAT Asn GTG ΙDΛ 766 7 гр CGG Arg CTC Leu 935 Trp **16**6 AGT Se r Polycystic Kidney Disease 999 110 Arg Phe ၁၉ ၁၉ CTA Leu 950 Pro AGC Ser CIC Leu ပ္ပ GTC Val 110 Phe 965 GTC cys 766 Trp ပ္ပ AIG Cys 885 ۷al 161 6TG Val 980 GTC Val CCG Pro 900 66A 61y AGC Se r ATC I le ၁၅၅ GIY ATG AAC Asn CCC Pro CAG <u>C</u> Met AAC ASN 915 ပ္ပ Pro CTG Leu CAG GIn GÁC CIG Asp GAA Glu GAG Glu 930 Leu GCA GCA Ald Val TTC Phe TCG Ser Phe GTA Val 945 GAG Glu GTA Val GTG D V ACC Thr ၅၁၅ AIG Arg 960 960 960 676 Val GTG Val **CGT** ACC Thr 880 CTG ပ္ပ ည္ဟ Ald Leu 975 Ald GCC A G ACG Thr GTG TCA Se r 895 ۷al GAG G I u TCC Ser GAG n S ASP 910 GTG TTC Phe GTG GAC Z Va ۷al CAG GIn GTG Val ၁၅ ပ္ပ Pro Len Leu Val



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FIG 3386 3434 3338 3242 3290 3146 3194 1100 1020 CAG GAG <u>n</u>9 GCC ACG Ala Thr 1035 <u>6</u> CAG GIn AId CAG Gin ပ္ပ AAC Asn **661** GI y Met TTC CCG GTT CCA GAC CCC TCG GTG GCC Phe Pro Val Pro Asp Pro Ser Val Ala ATG Glu Val GAG GTG His GIn Phe TCC AAC CAC GTG AGC Ser Asn His Val Ser GTC ATG CAC ACC TAC GCT GCC CCA Val Met His Thr Tyr Ala Ala Pro 1050 CAC CAG AGG Arg Asn AAT 1065 CGG ATG AAC / Arg Met Asn / 1015 GTG Ser Pro Leu Val Asp Ser Ala Val CTG TCC CCC 1080 Leu CIC ပ္ပ 90 000 V D I Q 1095 1CG Aig Vai Leu GGG GAG CAG (GI) / <u>n</u>19 1030 CTG GTG GAC GAG A d ACG GCC 1045 GTG Thr <u>|</u> 1060 ACC Thr SSS of ren CTG 1075 Asn Val 1090 TCC ACA GTG (Ser Thr Val I TCA Ser 995 AAC GTA Asn Vai GTG Asp Gly Val GGG GAT ACG GCG GGC Thr Ala Gly AAT Ser ည . 213 GI y 1010 ren GAG CAC / GAG 1025 Ш 0 lu Phe AAG TAC Tyr Lys Trp Thr 1 1055 TAC AAC Tyr Asn 1040 TGG ACC TTC Phe AAC ASN GTC Na! GTG Leu CTG ۷ď GIn CAG CTG ΙDΛ ΙDΛ Pro 1070 ည CTC CTC ren ACC Thr gç Y ren CIG Leu Ald CTG 1005 <u>G</u> Phe Za Va AId Leu GGT Val

3482

ASH GUC TIC GAG AAC CTG ACG CAG ASH ASH AIG Phe GIU ASH LEU Thr GIN

ACC GTG CTG GCA TCT Thr Val Leu Ala Ser

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1110

Applicant: Oxana Ibraghimov-Beskrovnaya, et al. Title: Compositions and Methods for Treating Polycystic Kidney Disease

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TAC

ACC TTC Thre The Teles

GTC ١

GCC GGC CGG CCC C Ala Gly Arg Pro V 1140

GAC GGC GTC CTG GTG Asp Gly Val Leu Val

AGT Ser

Val

1145

Tyr Pro

3530

GGT GLY

GCT GTG Ala Val

Ser Val

Pro ပ္ပ

ren CTG

SCC A d

AGC Ser

Pro Val

Val

CAG

ပ္ပ

Val Arg

1125 Ser

TCC

1130

12/59 3770 3626 3722 3674 Asn Asn Thr 1195 GAG GAG CTC Glu Glu Leu 1210 1180 ACC 999 Thr AAC AAC ACG <u>G y</u> His CAC Phe 110 ACG TGG GAC TThr Trp Asp P AAC Asn Pro Alg Alg A CAC GTG CGC CTG GAG GTC His Val Arg Leu Glu Val CCG GCT GCC Phe GCG GAT GTG CGC GTC T N AIG ASP VOI ARG VOI P 1205 1160 Val Arg Leu (1190 TAC CTT Ser Gin Pro Gly Gly Val Leu CAG AGC CAG **G11** Val Leu Thr Gin 1170 CCC CCT g Gly Thr Tyr Hi. 1185 ACC GCG GCG GCC CAG Ala Ala Ala GIn CIG CCT TCG Ser GTC AIG Ser Pro TCG AGG Arg Pro Leu Pro CCG CTG CCC TCC CCT <u>Gly</u> Ser 1150 AGC Ser ည္ဟ ည္ဟ AIG GI y ASP (1165 Tyr TAT



and the second			13/59			1T
	·					FIG. 1L
3818	3866	3914	3962	4010	4058	4106
0 Pro	G ACC	A GTG IF Val 1260	3T GCG y Ald 75	CTG GCC CGG AGC CTG CAC GTG CTG GTC TTC Leu Ala Arg Ser Leu His Val Leu Val Phe 1285	SG CAG	AC TAC
660 60 61y Al	ACG TG Thr Tr	GCA AC Alg Th	GTG GG Val G1 12	CTG G1 Leu VG 1290	CCC A(9 D T A I B
G CAG u GIn 1225	C ATC n Ile 40	G GAG O Glu	G ACC I Thr	c GTG s Val	C ATC S I le 1305	C CCG
6T6 GA Val G1	GAC AA Asp As	GGC CC Gly Pr 1255	CAT GTG TAC CTG CGG GCA CAG AAC TGC ACA GTG ACC GTG GGT GCG His Val Tyr Leu Arg Ala Gin Asn Cys Thr Val Thr Val Gly Ala 1275	CTG CA Leu Hi	GCC TG Ald Cy	GGG AA GLY AS 13
B GCC	3 GGC r G1y	S TCG	c TGC n Cys 1270	G AGC g Ser 85	0 A1a	C ACC
AGC CT(Ser Let 1220	SAG ACO	STG CT	CAG AA(SIn Asi	SCC CGAID AL	GAA CC Glu Pr 1300	TAC GT Tyr Vg
ATG /	GTG (ACC (Thr)	GCA (CTG	CTT (6CC A10 1315
rg GAC	00 000 DI	AC GGC SP G1y 125	re cec eu Arg 265	SC CAC	TG CGC eu Arç	TC ACC eu Thi
AGC GT Ser VG	AGC GC Ser Al	666 6/ 61 y A!	TAC C' Tyr Lo	AGC CCC GCC GGC CAC C Ser Pro Alg Gly His L 1280	616 C	CGG C
CTC Leu 1215	, GTC Val	ATG Met	6TG 8 Val	000 P	3 GAG u GIU 1295	c 6C6 p Ald 10
CGC GGA CTC AGC GTG GAC ATG AGC CTG GCC GTG GAG CAG GGC GCC CCC Arg Gly Leu Ser Val Asp Met Ser Leu Ala Val Glu Gln Gly Ala Pro 1215	GTG GTG GTC AGC GCC GCG GTG CAG ACG GGC GAC AAC ATC ACG TGG ACC Vai Vai Vai Ser Aia Aia Vai Gin Thr Giy Asp Asn Ile Thr Trp Thr 1230	TTC GAC ATG GGG GAC GGC ACC GTG CTG TCG GGC CCG GAG GCA ACA GTG Phe Asp Met Gly Asp Gly Thr Val Leu Ser Gly Pro Glu Ala Thr Val 1245	GAG CAT Glu His	GCC AGC Alg Sel	GTC CTG GAG GTG CTG CGC GTT GAA CCC GCC GCC TGC ATC CCC ACG CAG Vai Leu Giu Vai Leu Arg Vai Giu Pro Aia Aia Cys Iie Pro Thr Gin 1295	CCT GAC GCG CGC CTC ACG GCC TAC GTC ACC GGG AAC CCG GCC CAC TAC Pro Asp Ala Arg Leu Thr Ala Tyr Val Thr Gly Asn Pro Ala His Tyr 1310

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Kidney Disease

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						FIG.
4154	4202	4250	4298	4346	4394	4442
TGG ACC TTC GGG GAT GGC TCC TCC AAC ACG ACC GTG CGG Trp Thr Phe Gly Asp Gly Ser Ser Asn Thr Thr Val Arg 1330	ACG GTG ACA CAC ACC TTC ACG CGG AGC GGC ACG TTC CCC Thr Val Thr His Asn Phe Thr Arg Ser Gly Thr Phe Pro 1345	G CTG GTG CTG TCC AGC CGC GTG AAC AGG GCG CAT TAC TTC ACC a Leu Vai Leu Ser Ser Arg Vai Asn Arg Aia His Tyr Phe Thr 1360	ATC TGC GTG GAG CCA GAG GTG GGC AAC GTC ACC CTG CAG CCA GAG IIe Cys Vai Giu Pro Giu Vai Giy Asn Vai Thr Leu Gin Pro Giu 1375	GTG CAG CTC GGG GAC GAG GCC TGG CTG GTG GCA TGT GCC Vai Gin Leu Gly Asp Giu Aia Trp Leu Vai Aia Cys Aia 1395	TTC CCC TAC CGC TAC ACC TGG GAC TTT GGC ACC GAG GAA Phe Pro Tyr Arg Tyr Thr Trp Asp Phe Gly Thr Glu Glu 1410	TCC CGT GCC AGG GGC CCT GAG GTG ACG TTC ATC TAC CGA Thr Arg Ala Arg Gly Pro Glu Val Thr Phe Ile Tyr Arg 1425
CTC TTC GAC TGG ACC T Leu Phe Asp Trp Thr P 1325	GGG TGC CCG A	CTG GCG CTG Leu Aig Leu	AGC ATC TGC Ser IIe Cys	AGG CAG TTT GTG (Arg Gin Phe Val (1390	TGG CCC CCG Trp Pro 1405	GCC GCC CCC Ald Ald Pro

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·						FIG.
4490	4538	4586	4634	4682	4730	4778
Ser	GTC Val	CAG CCG GIn Pro	GTG GGC CGT GGG CGC CCC GCC AGC TAC CTG TGG Val Gly Arg Gly Arg Pro Ala Ser Tyr Leu Trp 1490	GCT	GAG G1u	616 Va I
ATC Ile	CTG	CAG GIN	CTG	CAC His 1515	AAT Asn	CGC Arg
AAC ASn ASn 1450	GTG	CAG GIn	TAC Tyr	ACC Thr	166 1 r p 1530	CGG Arg
CTT GTG ACA GTC ACC GCG TCC AAC AAC ATC TCT Leu Val Thr Val Thr Ala Ser Asn Asn Ile Ser 1445	GCC CTG GTG GAG GAG CCC GTG CTG GTC Ala Leu Val Glu Val Gln Glu Pro Val Leu Val 1460	AAT GGC TCC CTT GGG CTG GAG CTG CAG Asn Gly Ser Leu Gly Leu Glu Leu Gln 1475	AGC	GGG TGG CTC GAG GGT CCG GAG GTC ACC CAC GCT Gly Trp Leu Glu Gly Pro Glu Val Thr His Ala 1510	GAC TTC ACC GTT AGG GTG GCC GGC TGG AAT GAG Asp Phe Thr Val Arg Val Ala Gly Trp Asn Glu 1525	GCC TGG CTC AAT GTG ACG GTG AAG CGG CGC GTG Ala Trp Leu Asn Val Thr Val Lys Arg Arg Val 1540
TCC	o Ac Clu	GAG G I U 1480	D I A	GAG Glu	D I V	GTG Val
GCG A I a	CAG	CTG	CCC Pro 1495	CCG Pro	GTG Val	ACG Thr
ACC Thr	GTG Val	666 617	CGC Arg	GGT G1 y 1510	AGG Arg	616 Val
GTC Val 1445	GAG G I u	CTT	666 61 y	GAG GIU	GTT Val 1525	AAT Asn
ACA Thr	GTG Val 1460	Ser	CGT Arg	CTC	ACC Thr	CTC Leu 1540
GTG Val	CTG	660 61 y 1475	000 01y	766 7rp	TTC Phe	166 1rp
CTT	BIA	AAT Asn	GTG Val 1490	666 617	GAC	B W
TAT	Ser	GTC	GCT	66T 61y 1505	66T 61y	CAG G I u
TCC Se r 1440	GAC	AAG Lys	TCT Ser	GAC GGT G Asp Gly G	AGC ACA GGT Ser Thr Gly 1520	AGC Ser
66C 61 y	AAT Asn 1455	ATC Ile	TTC Phe	999 61 ×	AGC Ser	CGC Arg 1535
CCA GGC TCC TAT Pro Gly Ser Tyr 1440	GCC AAT GAC Alg Asn Asp 1455	AGC Ser 1470	CTG	CTG	AAC Asn	AGC CGC AGC GAG Ser Arg Ser Glu 1535
GAC Asp	GCT	ACC AGC ATC AAG Thr Ser IIe Lys 1470	TAC CTG TTC TCT Tyr Leu Phe Ser 1485	GAT Asp	TAC	GTG Vai



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n.			16/59			10
·						FIG. 10
4826	4874	4922	4970	5018	5066	5114
GCA AGC CGC ACG GTG GTG CCC CTG AAT GGG Aig Ser Arg Thr Vai Vai Pro Leu Asn Gly 1555	TCG CTG GAG GCC GGC AGT GAT GTG CGC TAT Ser Leu Giu Aig Gly Ser Asp Vai Arg Tyr 1575	CGC TGC ACG CCC ATC CCT GGG GGT CCT ACC Arg Cys Thr Pro Ile Pro Gly Gly Pro Thr 1590	TCC GTG GGC ACC TTC AAT ATC ATC GTC ACG Ser Val Gly Thr Phe Asn Ile Ile Val Thr 1605	TCC GCC CAG GAC AGC ATC TTC GTC TAT GTC Ser Ala Gin Asp Ser Ile Phe Val Tyr Val 1620	CTG CAG GTG GGC GGT GGC CGC TAC TTC Leu Gin Vai Vai Gly Gly Giy Arg Tyr Phe 1635	CAG CTG CAG GCC GTG GTT AGG GAT GGC ACC Gin Leu Gin Aia Vai Vai Arg Asp Gly Thr 1655
CGG GGG CTC GTC GTC AAT Arg Gly Leu Val Val Asn 1550	AGC GTG AGC TTC AGC ACG TO Ser Val Ser Phe Ser Thr Ser 1565	TCC TGG GTG CTC TGT GAC Ser Trp Val Leu Cys Asp 1585	ATC TCT TAC ACC TTC CGC TILE Ser Tyr Thr Phe Arg S	GCT GAG AAC GAG GTG GGC Ala Glu Asn Glu Val Gly 1615	CTG CAG CTC ATA GAG GGG Leu Gin Leu Ile Giu Gly 1630	CCC ACC AAC CAC ACG GTA Pro Thr Asn His Thr Val 1645



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			17/59			1P
		,				FIG. 1P
5162	5210	5258	5306	5354	5402	5450
AAC GTC TCC TAC AGC TGG ACT GCC TGG AGG GAC AGG GGC CCG GCC CTG Asn Val Ser Tyr Ser Trp Thr Ald Trp Arg Asp Arg Gly Pro Ald Leu 1675	GGC AGC GGC AAA GGC TTC TCG CTC ACC GTG CTC GAG GCC GGC ACC Gly Ser Gly Lys Gly Phe Ser Leu Thr Val Leu Glu Ala Gly Thr 1680	C CAT GTG CAG CTG CGG GCC ACC AAC ATG CTG GGC AGC GCC TGG GCC r His Val Gin Leu Arg Aig Thr Asn Met Leu Giy Ser Aig Trp Aig 1705	GAC TGC ACC ATG GAC TTC GTG GAG CCT GTG GGG TGG CTG ATG GTG GCC Asp cys thr Met Asp Phe Vai Giu Pro Vai Giy Trp Leu Met Vai Aid 1710	GCC TCC CCG AAC CCA GCT GCC GTC AAC ACA AGC GTC ACC CTC AGT GCC Alg Ser Pro Asn Pro Alg Alg Val Asn Thr Ser Vgl Thr Leu Ser Alg 1725	CTG GCT GGT GGC AGT GGT GTC GTA TAC ACT TGG TCC TTG GAG GAG Leu Aia Giy Giy Ser Giy Vai Vai Tyr Thr Trp Ser Leu Giu Giu 1745	GGG CTG AGC TGG GAG ACC TCC GAG CCA TTT ACC ACC CAT AGC TTC CCC GIY Leu Ser Trp Giu Thr Ser Giu Pro Phe Thr Thr His Ser Phe Pro 1760 1760
AAC	DIA PIQ	TAC Tyr	GAC Asp	GCC A10 1725	GAG	666 61 <i>y</i>



C48

5498		
၁၅၅	Leu Gly	
CTG	ren	-
900	Pro	
AAC	Asn Pro	1785
999		•
GCA	Aid	
ACG	Thr	
ATG	Met	_
ACC	Thr N	1780
GTC	D/	
116	Leu	
CAC	His	
CTG	Leu	
၁၅၅	Gly Leu	1775
ည္ဟ	Pro	

ACA Thr

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5546 ၁၅၅ AGT Ser GTG GAA GTG GAT GTG CAG GTG CCT GTG Val Glu Val Asp Val Gln Val Pro Val 1800 Asp Val ACC Thr GCC AAC GCC A 1790 TCA

5594 1820 666 GI y Ald ည္ဟ OCG A d TTC GTG Phe Val GGA GGC AGC TO GIY GIY Ser P 1815 AGC GAG CCC (Ser Glu Pro (1810 GCC A I a AGG Ile Arg ATC Leu Ser AGC

5642 Asn Val Ser 1835 AGC AAT GTG CAG CTG GCC ACG GGC ACC GIn Leu Aig Thr Gly Thr 1830 Trp Gly 166 666 Phe 1825 111 Prol ပ္ပ GTG Va Va 101 Ser ည Ser

5690 GTC ΔV Arg Gly Pro His CCT CAT AAG CGT GGC (Lys Arg Gly F GGC AGC AGC A Gly Ser Ser 1 1845 299 222 Pro Gly TGG GCT GTG C Trp Ala Val P 1840 **1**60 Cys 166 1 r p

5738 TCC ATC CGG CTC AAT GCC Ser Ile Arg Leu Asn Ald Arg Leu Asn i 1865 GCT GGC ACC TTC T Ala Gly Thr Phe S 1860 CCG GAT GCT (Asp GTC TTC (Val Phe I Val 1855 ATG Met ACC Thr

5786 GAG GTu GTC TCA GCC ACG TAC AAC CTC ACG GCG Vai Ser Aid Thr Tyr Asn Leu Thr Aid Val 1875 <u>9</u>91 AAC GCA GTC AGC ASN AIG VOI Ser 1870 ည Ser

FIG. 10

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			19/59			1R
						FIG. 1R
5834	5882	5930	5978	6026	6074	6122
GAG CCC ATC GTG GGC CTG GTG CTG TGG GCC AGC AGG GTG GTG GCG Glu Pro Ile Val Gly Leu Val Leu Trp Ala Ser Ser Lys Val Val Ala 1885 1890	GGG CAG CTG GTC CAT TTT CAG ATC CTG CTG GCT GCC GGC TCA GCT GIN Leu Val His Phe GIn Ile Leu Leu Ala Ala GIY Ser Ala 1905	S ACC TTC CGC CTG CAG GTC GGC GGC GAC CCC GAG GTG CTC CCC ITC CCC ITC CTC CCC ITC The Arg Leu Gin Vai Giy Giy Aia Asn Pro Giu Vai Leu Pro 1920	s CCC CGT TTC TCC CAC AGC TTC CCC CGC GTC GGA GAC CAC GTG GTG y Pro Arg Phe Ser His Ser Phe Pro Arg Val Gly Asp His Val Val 1935	AGC GTG CGG GGC AAA AAC CAC GTG AGC TGG GCC CAG GCG CAG GTG CGC Ser Val Arg Gly Lys Asn His Val Ser Trp Ala Gln Ala Gln Val Arg 1950	ATC GTG GTG CTG GAG GCC GTG AGT GGG CTG CAG GTG CCC AAC TGC TGC Ile Val Val Leu Glu Ala Val Ser Gly Leu Gln Val Pro Asn Cys Cys 1965	GAG CCT GGC ATC GCC ACG GGC ACT GAG AGG AAC TTC ACA GCC CGC GTG Giu Pro Giy Ile Aig Thr Giy Thr Giu Arg Asn Phe Thr Aig Arg Vai 1985 1990
GAG G1u 1885	0 0 0 1	GTC V01	666 619	AG Se	AT II.	8 9 -







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						FIG
6170	6218	9979	6314	6362	6410	6458
CGG GTC GCC TAC TGC TTC TCG CTG CAG AAG Arg Val Ala Tyr Ala Trp Tyr Phe Ser Leu Gin Lys 2005	TCG CTG GTC ATC CTG TCG GGC CGC GAC GTC ACC TAC Ser Leu Val Ile Leu Ser Gly Arg Asp Val Thr Tyr 2020	GCG GGG CTG TTG GAG ATC CAG GTG CGC GCC TTC AAC Ala Gly Leu Leu Glu Ile Gln Val Arg Ala Phe Asn 2035	TT CAG GAC GCC ai Gin Asp Ala 2060	GCC CTG CAG AGC GGC CCC TGC TTC ACC AAC CGC TCG Ala Leu Gin Ser Giy Pro Cys Phe Thr Asn Arg Ser 2065	GCC GCC ACC AGC CCC CGG CGT GTG GCC TAC Ala Ala Thr Ser Pro Ser Pro Arg Arg Val Ala Tyr 2090	GGG GAT GGG TCG CCA GGG CAG GAC ACA GAT GAG CCC Gly Asp Gly Ser Pro Gly Gln Asp Thr Asp Glu Pro 2100
IGG TAC TTC TC Irp Tyr Phe Se	TCG GGC CGC G/ Ser Gly Arg A:	ATC CAG GTG CO Ile Gin Val Ai 2040	GAG AAC CGC ACG CTG GTG CTG GAG GTT CAG GAC GIU ASn Arg Thr Leu Vai Leu Giu Vai Gin Asp 2050	CCC TGC TTC AO Pro Cys Phe Ti 2070	AGC CCC CGG C	GGG CAG GAC A Gly Gin Asp T
CC TAC GCC 1	TC ATC CTG 1	TG TTG GAG / eu Leu Glu] 035	SC ACG CTG (AG AGC GGC (CC AGC CCC /	36G TCG CCA (31y Ser Pro 2100
T CGG GTC G r Arg Val A 00	c TCG CTG G p Ser Leu V				G GCC GCC / u Ala Ala 1 80	T GGG GAT (le Gly Asp (
CAG CGC GGC TCT GIN Arg Gly Ser	GTC CAG GGC GAC Vai Gin Gly Asp 2015	ACG CCC GTG GCC Thr Pro Val Ala 2030	GCC CTG GGC AGT Ald Leu Gly Ser 2045	GTC CAG TAT GTG Vai Gin Tyr Vai	GCG CAG TTT GAG Ala Gin Phe Glu 2080	CAC TGG GAC TTT His Trp Asp Phe 2095
ပ်ပ	Θ >	Ā⊢	N A N	S >	ຜ ∢	OI

FIG. 1S

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6506		
STG :	ΙDΛ	
CAG	GIn	
CTG CAG	Val	•
ည္ဟ	Arg	_
TAC CGC	Tyr	2120
GAC	Asp	
999	Gly Asp Tyr Arg	
CCT	J Pro	
AGG	Arg	
CTG	Leu	2115
TAC	Tyr	
CAC TCC	Ser	
CAC	His Ser Tyr Leu Arg	
GAG	<u>n</u> 19	_
ည္ဟ	Ald	2110
AGG GCC (Arg Ald	•

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6554		
ACC	Thr	2140
GTG	<u>D</u>	
ACG	Thr	
ည္ဟ	AIG	
CAG	GIn	
939	AIG	2135
6TG	Vai Ala (
110	Phe	
110	Phe	
AGC	Ser	_
GTG	<u> </u>	2130
CTG	Leu	
AAC	Asn	
100	Ser	
သ	Asn Ald	
AAC	ASN	2125

6602		
STC CTG CCC	Pro	
CTG	Leu	2155
GTC	<u> </u>	
GTG	<u>p</u>	
	ASp	•
GTG	<u>D</u>	
GAG	<u>n</u> 19	2150
ည္ဟ	Pro	
GAG (<u>n</u> 9	
999	Arg	•
1 60	Cys	
ည္ဟ	Ald Cys	2145
CTG	Val Leu	
616	ΙDΛ	
GTC CAG (GIn	
GTC	۲d	

6650			6698
CAC	His		GAG
ည္ဟ	A I d		16G
GAG	n l o	21/0	ည္ပ
116	ren		TAC
TAC	g Ser Gin Arg Asn Tyr Leu Giu Ala H		GAG TAC
AAC	Asn		CAG ACT
ည္ဟ	Arg	_	CAG
CAG	Gin	2165	TAC
TCA	Ser		GTC ACC
CGA	Arg		3 CTC
၁၅	Arg		291
ATG	Met	_	GAC
CTG	Leu	716(၁၅၁
616	Val		CTG
CAG	Leu Gin Vai Leu Met Arg Arg		GAC
CTG	ren		GTT

8699		
GAG	n 19	
166	Trp	•
၁ဗ၁	Tyr Arg	
TAC	Tyr	2185
GAG	n Cln	
ACT	Thr	
CAG	Gin	
TAC	Tyr	_
ACC	Thr	2180
GTC	Val	
	Cys	•
GAC	Asp	•
၁၅	Leu Arg	
CTG	Leu	2175
	ASp	•
GTT	Va!	

6746		
GTG	n Arg Pro Gly Arg Pro Ala Arg Val	
CGT	Arg	
939	Ald	
SS	Pro	
ည	Arg	2200
999	<u>cl y</u>	
၁	Pro	
993	Arg	
CAG	<u>el</u> n	
160	Cys	2195
AGC	a Ser Cys Gln /	
ည္ဟ	r Arg Thr Ala	
ACC	Thr	
ည္ဟ	Arg	_
TAT	I Tyr A	2190
GTG	Va.	

	FIG
6794	•
993	Arg 2220
၁	Pro
STG CTG CCG	ren
GTG	ID/
CTG	Leu
990	Arg Pro Arg Leu ' 2215
CCT	Pro
990	Arg
AGC	Ser
, 676	D _
GAC	ASP 2210
GTG	ID/
၁၅၅	G y
ည္ဟ	Pro
CTG	Leu
SCC CTG CCC GGC	A10 2205



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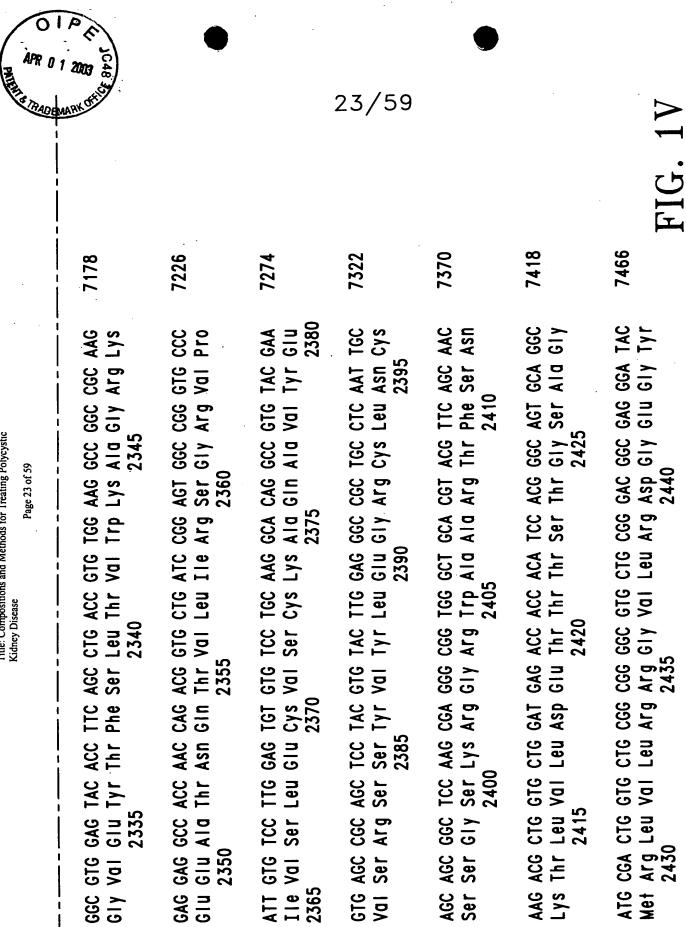
	•		•	٠		
						FIG. 11
6842	6890	6938	6986	7034	7082	7130
CAC TAC TGC TTT GTG TTT GTC GTG TCA TTT His Tyr Cys Phe Val Phe Val Val Ser Phe 2230	T GTG ACG GTG GCC n Vai Thr Vai Ala 2250	G 166 I Trp	TCA GAC ACA CGG GAC CTG GTG CTG GAT GGG AGC GAG TCC TAC GAC CCC Ser Asp Thr Arg Asp Leu Val Leu Asp Gly Ser Glu Ser Tyr. Asp Pro 2270	C TGT a Cys 2300	GGC GGG TGT GCG CTG AAC TTT GGG Gly Gly Cys Ala Leu Asn Phe Gly 2310	CGC GGG AGC AGC ACG GTC ACC ATT CCA CGG GAG CGG CTG GCG GCT Arg Gly Ser Ser Thr Val Thr Ile Pro Arg Glu Arg Leu Ala Ald 2320
Ser 2235	DA .) CT(GA(9 P C	C TT n Ph 23	G GC U A1
GTG Val	ACG Thr 225	CGC Arç	TAC Tyl	166	AA(CT()
GTC Val	GTG Val	TAC Tyr 226	TCC Ser	CAC His	CTG Leu	CGG
TTT Phe	AAT Asn	TCA TAC CGC GTG T Ser Tyr Arg Val T 2265	GAG G1u 228(TTC Phe	B I A	GAG G I u
GTG Val	BIA	000 01 y	AGC Ser	AGT Ser 2295	16T Cys	CGG Arg
TTT Phe 2230	CAG	. GGT GGC .	666 61 y	CTC	666 61 y 2310	CCA Pro
16C Cys	AGC ATC CAG GCC AAT Ser IIe Gin Aig Asn 2245	GAG GIU	GAT ASp	CCG Pro	66C 61y	ATT 11e 232
TAC Ty r	AGC Ser	ATT Ile 2260	CTG Leu	ACG Thr	GCT	ACC Thr
CAC His	SAG	ATC Ile	GTG Val 2275	CAG GIN	0A6 Glu	GTC Val
666 61y	ACA Thr	CCC Pro	CTG	GAC ASP 229(AGG Arg	ACG Thr
CTG CCT GTG GGG Leu Pro Val Gly 2225	C ACG CCA CTG ACA (P Thr Pro Leu Thr C 2240	GTG CCC ATC ATT GAG Val Pro Ile Ile Glu 2260	GAC Asp	GGC GAC CAG ACG CCG CTC AGT TTC CAC TGG GCC Gly Asp Gln Thr Pro Leu Ser Phe His Trp Ald 2290	GCT TCG ACA CAG AGG GAG GCT Ala Ser Thr Gin Arg Giu Ala 2305	AGC Ser
CCT Pro	CCA Pro 2240	CTG	CGG Arg	GAC	ACA Thr	AGC Ser 232
CTG	ACG Thr	CGC Arg 2255	ACA Thr	GAG Glu	TCG Ser	666 61y
B I A	GA(AS	GAG CGC CTG Glu Arg Leu 2255	GAC ASP 2270	CTG	GCT	CGC Arg
CTG	666 619	CCC Pro	TCA	AAC CTG GAG GAC (Asn Leu Glu Asp (2285	GTG Val	0 0 0 0

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GAG

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AAG Lys

Met

AGC Se r

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7514		
160	Cys	2460
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GAG	<u>n</u> 19	
GAG (<u>n</u> 9	
GAG	n 9	
၁၅၅	<u>Gl</u>	2455
TCT		
၁၅၁	Arg	
၁၅၅	7 61	•
CTG	Leu	
GTG	Val Leu	2450
ACG	Thr	
CTC	Leu	
ACG	Thr	
TTC	Thr Phe	
ACC	Thr	2445

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7562 ATC CGC CTG TCC CCC AAC CGC CCG CCG CTG GGG GGC TCT TGC Ile Arg Leu Ser Pro Asn Arg Pro Pro Leu Gly Gly Ser Cys 2475 2470 2465 TCC ည္ဟ Ald

7610 CAC His AAG GTG (Lys Val H 2490 Gly Ala Val His Ala Leu Thr Thr 2485 GTG CAC GCC CTC ACC ACC SCT ည္တ CCA CTG Phe Pro Leu 110 ren CIC Arg

7658 CTG Asp Ald Gly Ald Pro Leu 2505 900 009 099 GAT GCT Asp Ala Glu A 2500 GAC GCG GAG u Cys Thr Gly Trp H:-2495 GAA <u>n</u>19 Phe

7706 TGC GAG GAG Cys Glu Glu GIN GIY HIS (2520 CAG GGC CAC GCC CTG CTG CTG CGC CGC TGT CGC Leu Arg Arg Cys Arg 2515 Tyr Ala Leu Leu 2510 TAC Val

7754 : GTG CTG CCC I Val Leu Pro 2540 TAC GGA GCC (Tyr Gly Ald V 2535 GTC TAC AAG GGC AGC CTC TCC AGC Val Tyr Lys Gly Ser Leu Ser Ser 2530 Phe Cys 2525 161

7802 GTG CAG Val GIN 2555 GTG Val GGC CTG GCC GTG ۲a AId Gly Leu / 2550 CTG (D/ TTC GAG (Phe Glu CCA CAC 1 Pro His F 2545 TTC AGG Arg Phe <u>CI y</u> **GGT** ပ္ပ Pro

FIG. 1W



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				•		
						FIG.
7850	7898	7946	7994	8042	8090	8138
CAG CTG GGA GCC GCT GTG GTC GCC CTC AAC AGG TCT TTG GCC ATC Gin Leu Giy Aid Aid Vai Vai Aid Leu Asn Arg Ser Leu Aid Ile 2560	GGG CTC ACA GTC TGG CTG Gly Leu Thr Val Trp Leu 2585	CAC GGG CTC ACC GCT AGT GTG CTC CCA GGG CTG CTG CGG CAG GCC GAT His Gly Leu Thr Ala Ser Val Leu Pro Gly Leu Leu Arg Gln Ala Asp 2590	AAC ASN 2620	CGG GCC CTG GAC GTG GCG GCA GAG CCC AAG CAC GAG CGG Arg Aig Leu Asp Val Aig Aig Giu Pro Lys His Giu Arg 2625	CAC CGA GCC CAG ATA CGC AAG AAC ATC ACG GAG ACT CTG GTG TCC His Arg Ala Gin Ile Arg Lys Asn Ile Thr Giu Thr Leu Val Ser 2640	CTG AGG GTC CAC ACT GTG GAT GAC ATC CAG CAG ATC GCT GCT GCG CTG Leu Arg Val His Thr Val Asp Asp Ile Gin Gin Ile Ala Ala Ala Leu 2655
000 V V	166 Trp	SCC A I d	CTG Leu	GAG G11 263	610 Val	9 A
TTG Leu 2570	61C Val	CAG GIn	GTG Val	CAC His	CTG Leu 265	GCT Ald 5
TCT Ser	ACA Thr 2585	CGG Arg	ACC Thr	AAG	ACT Thr	GCT A I d 266
AGG Arg	CTC	CTG Leu 260(GTC Val	CCC Pro	GAG G1u	ATC Ile
AAC Asn	666 61 y	CTG	CTG Leu 2615	oAc GIU	ACG Thr	CAG GIn
CTC	ACG	666 61 y	GCC	6CA A10 2630	ATC I le	CAG
GCC A I a 2565	CCC AAC GGC AGC GCA ACG Pro Asn Gly Ser Ald Thr 2580	CCA Pro	TTG Leu	B I A	AAC ASN 264	ATC Ile
GTC Val	AGC Ser 2580	CTC Leu	TCG Ser	6T6 Val	AAG Lys	GAC ASP 266(
GTG Val	660 61	GTG Val 2595	TAC	GAC	CGC	GAT Asp
GCT	AAC Asn	AGT Ser	GAG G I U 2610	CTG Leu	ATA I le	GTG Val
BIA (000 Pro	GCT	ATC I le	GCC A10 2625	CAG GIN	ACT Thr
GGA G1y 2560	CTC CCA GAG Leu Pro Glu 2575	ACC Thr	GTC Val	CGG Arg	GCC A10 264	CAC His
CTG Leu	CCA Pro 2575	CTC	CAC His	TAC GAG Tyr Glu	CGA Arg	GTC Val 265
CAG	CTC	666 61 y 2590	CAG	TAC Tyr	CAC His	AGG Arg
GAC	ACC Thr	CAC His	CCC CAG CAC GTC ATC GAG TAC TCG TTG GCC CTG GTC ACC GTG CTG Pro GIn His Val IIe Glu Tyr Ser Leu Ala Leu Val Thr Val Leu 2605	GAG Glu	CAG GIn	CTG
					•	

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8186 CTG Cys Arg Ser Cys 16C CGC 2680 Gly Pro Ser Arg Glu Leu Val GAG AGC AGG 2675 ပ္ပ 999 Met GIn Cys ည 2670

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8234 2700 CAG GCA Gin Aid CAG CTG Met Leu Ile Leu ATC CIC 2692 ATG ATG Aid Met ပ္ပ GAG 01n Lys Leu (2690 AAG CTG His CAC Leu CTG Thr ACG Lys GIn . 2685 CAG

8282 Asp Ser Ile 2715 ATC AGC GAC GGA <u>G</u> ATC Thr Ald ACC GCC 2710 . 6 2 ပ္ပ ACG Thr GGC ACC GTG / GIY Thr Val 1 2705 AIG ည္ဟ Thr ACC Thr ACC GAG

8330 Arg 999 Asp Val . 2730 GTG GAC Ser Ser ATC CAC CTG GCC AGC TCG Ald IIe His Leu / 2725 GAC CTC Asp Leu ACA GGA (Thr Gly Ile ATC AAC Asn Leu

8378 GTG Va TCA CCA TCT CGG ATG Ser Pro Ser Arg Met Pro Ser Arg N 2745 Gly Ald Glu : 2740 GGA GCC GAG CIG ren GAG n 19 TCA Ser ပ္ပ GIN Pro CAG CCA **GCA**

8426 ATG Met CTC ATG CGC ATC CTC A Leu Met Arg IIe Leu M 2760 TCT GCC Ser Ald CTG ACC | Leu Thr 9 2755 Asn AAC TAC Tyr ပ္ပ AId CAG G G I Ser ည

8474 2780 GAG <u>n 19</u> GAG 960 610 GAG CCC CTG ACG CTG GCG Glu Pro Leu Thr Leu Ald 2775 AAC GAG Asn Glu CTC Leu 000 CGC Nal Arg Arg Ser 2765 ည ည ် ပုပ္ပ

FIG.

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AF AF	R 0 1 2003 &	2
8522	8570	8618
CCG CGG AGC CTG CTG TGC TAT Pro Arg Ser Leu Leu Cys Tyr 2790	CCT GGC TGC CAC TTC TCC ATC CCC GAG GCT TTC Pro Gly Cys His Phe Ser Ile Pro Glu Ala Phe 2805	G CTC ATC TTT CTG n Leu IIe Phe Leu 2825
GAC CCG CGG AG R ASP Pro Arg Se 2790	CAC TTC TCC ATS His Phe Ser II	AAC CTC AGT GAC GTG GTG CAG CTC ATC TTT Asn Leu Ser Asp Val Val Gin Leu Ile Phe 2820
CAG GGC AAG CGC TCG GAC GIN GIY LYS Arg Ser Asp 2785	GGG CCT GGC TGC Gly Pro Gly Cys	GCC AAC CTC AGT Ald Asn Leu Ser 282
GCC CAG Ala Gin	GCC CCA GGG Ald Pro Gly 2800	GCC CTG GCC Alg Leu Alg 2815

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GTG Ile Val 999 GI y

AGC Ser

9998	8714
GTC Val	CAG G1n 2860
ACC Thr	B I A
TAC Ty r	66C 61 y
AAC Asn	B I A
AGC Ser 2840	CAG
ATC Ile	ACA Thr 2855
TAT Tyr	CAG GIn
66C 61 y	TTC Phe
TTT Phe	GCA A I a
CCC Pro 2835	ATG Met
TTT Phe	TCG / Ser 2850
CCC Pro	B I A
AAT Asn	GTG Val
Ser	AAG Lys
GAC ASP 2830	ACC
GTG Val	TCC Ser 2845

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G AAG GTG	II Lys vai 2875	
C ACC GT	e ihr va	
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CA GAG CG	er Giu Ar 28	
TG GCC T(Glu Arg Leu Ald Ser Glu Arg Ald Ile Ihr Val 2865	
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AAC		
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100	r Ser Ald	2890
AG	Se	
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990	Arg	2885
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GCT	A I d	
166	Trp Ald	•
GAC	Asp	
100	Ser Asp	2880
AAC	Asn	
AAC	Asn	
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Ser

2910

Val

Tyr

Asn 2925

TAT

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28/59 FIG. 1AA 9146 9098 9050 9002 8858 8906 8954 2940 CTG GTG , GAG CCC Glu Pro AIG Asp AGA GAC CCA GCG Leu GAG CAC AAC GGT GCT GAC CTG CAT CTG CAG CTC Leu His Leu GIn Leu Glu His Asn CGC TGG TCG GCG C Arg Trp Ser Ald L 3000 Arg Asp Pro . 2985 GIY AID 2970 2955 Ser Val Gly Ala င္သ Ser Glu Glu Pro GTC GGT 2905 CGC CCA GAG TCA CTC CAG Arg Pro Glu Ser Leu Gin 2965 TCT GAG GAA Glu Pro Arg Pro Asn CGG CCC AAT AGC Ser 2920 ည TTC 2935 666 GI y AGC AAC CCT GCG GCC GGG Ser Asn Pro Ala Ala Gly Gin Pro Gin Ala 2950 TTC ATT TCC CCG Phe IIe Ser Pro CTC TCC AGC CAC Leu Ser Ser His CAG CCC CTG Tyr Leu Ser TAC CAG CCC u Asp Gly His Tv. 2930 2980 2900 **1**00 Ser s GCT AGC AGG AGG ATC C r Ala Ser Arg Arg Ile A 2960 2915 Tyr Leu His 2945 TCC GTT GTG GTC Ser Val Val Val TAC CTA CAC TTC Phe TAC CAT CTG AAC Asn CGG CCC TAC ACC T Arg Pro Tyr Thr P 2975 ACC CTG GAC AGC / Thr Leu Asp Ser S Leu CIG Thr Leu Leu ID/ His CIG GTC Ser Tyr 2895 ACG QC V D A Ald Asn

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Ser

Cys

CAC

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TAC Tyr AGT

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TRADEMA	RK OTHER		29/59			1AB
9194	9242	9290	9338	9386	9434	9482 FIG. 1AB
CAG GTG TCC GTG GGC CTG TAC ACG TCC CTG TGC CAG TAC TTC AGC GAG GIN Val Ser Val Gly Leu Tyr Thr Ser Leu Cys Gin Tyr Phe Ser Glu 3005	GAG GAC ATG GTG TGG CGG ACA GAG GGG CTG CTG CCC CTG GAG GAG ACC Glu Asp Met Val Trp Arg Thr Glu Gly Leu Leu Pro Leu Glu Glu Thr 3025	TCG CCC CGC CAG GCC GTC TGC CTC ACC CGC CAC CTC ACC GCC TTC GGC Ser Pro Arg Gin Ala Val Cys Leu Thr Arg His Leu Thr Ala Phe Gly 3040	GCC AGC CTC TTC GTG CCC CCA AGC CAT GTC CGC TTT GTG TTT CCT GAG Alg Ser Leu Phe Val Pro Pro Ser His Val Arg Phe Val Phe Pro Glu 3055	CCG ACA GCG GAT GTA AAC TAC ATC GTC ATG CTG ACA TGT GCT GTG TGC Pro Thr Ala Asp Val Asn Tyr Ile Val Met Leu Thr Cys Ala Val Cys 3070	CTG GTG ACC TAC ATG GTC ATG GCC GCC ATC CTG CAC AAG CTG GAC CAG Leu Vai Thr Tyr Met Vai Met Aia Aia Iie Leu His Lys Leu Asp Gin 3085 3100	TTG GAT GCC AGC CGC GCC ATC CCT TTC TGT GGG CAG CGG GGC Leu Asp Ala Ser Arg Gly Arg Ala Ile Pro Phe Cys Gly Gln Arg Gly 3115

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APR 0 1 200	10.48 JULY		30/59			FIG. 1AC
9530	9578	9626	9674	9722	9770	9818 FIG
TTC AAG TAC GAG ATC CTC GTC AAG ACA GGC TGG GGC CGG GGC TCA Phe Lys Tyr Glu Ile Leu Val Lys Thr Gly Trp Gly Arg Gly Ser 3120	ACC ACG GCC CAC GTG GGC ATC ATG CTG TAT GGG GTG GAC AGC CGG Thr Thr Ala His Val Gly Ile Met Leu Tyr Gly Val Asp Ser Arg 3135	AGC GGC CAC CGG CAC CTG GAC GGC GAC AGA GCC TTC CAC CGC AAC AGC Ser Gly His Arg His Leu Asp Gly Asp Arg Ald Phe His Arg Asn Ser 3150	CTG GAC ATC TTC CGG ATC GCC ACC CCG CAC AGC CTG GGT AGC GTG TGG Leu Asp Ile Phe Arg Ile Alg Thr Pro His Ser Leu Gly Ser Val Trp 3165	AAG ATC CGA GTG TGG CAC GAC AAC AAA GGG CTC AGC CCT GCC TGG TTC Lys Ile Arg Val Trp His Asp Asn Lys Gly Leu Ser Pro Ala Trp Phe 3185	CAG CAC GTC ATC GTC AGG GAC CTG CAG ACG GCA CGC AGC GCC TTC GIN His Val Ile Val Arg Asp Leu Gln Thr Ala Arg Ser Ala Phe 3200	G GTC AAT GAC TGG CTT TCG GTG GAG ACG GAG GCC AAC GGG GGC u Val Asn Asp Trp Leu Ser Val Glu Thr Glu Ala Asn Gly Gly 3215 3220 3220
CGC TAPP PI	GGT A0	AGC G Ser G	CTG G Leu A 3165	AAG A Lys I	CTG CAG Leu GIn	TTC CTG Phe Leu

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APR 0 1 2003	()		31/59			FIG. 1AD
9866	9914	3962	10010	10058	10106	10154 FIG
GCC GCG AG Ala Ala Se 5	rg ctg gtg gct gag ctg cag cgt ggc ttc ttt gac Aag su Leu Val Ala Glu Leu Gln Arg Gly Phe Phe Asp Lys 3250	CAC ATC TGG CTC TCC ATA TGG GAC CGG CCG CCT CGT AGC CGT TTC ACT His Ile Trp Leu Ser Ile Trp Asp Arg Pro Pro Arg Ser Arg Phe Thr 3275	CGC ATC CAG AGG GCC ACC TGC TGC GTT CTC CTC ATC TGC CTC TTC CTG Arg Ile Gin Arg Aid Thr Cys Cys Vai Leu Leu Ile Cys Leu Phe Leu 3280	GGC GCC AAC GCC GTG TGG TAC GGG GCT GTT GGC GAC TCT GCC TAC AGC Gly Ala Asn Ala Val Trp Tyr Gly Ala Val Gly Asp Ser Ala Tyr Ser 3295	ACG GGG CAT GTG TCC AGG CTG AGC CCG CTG AGC GTC GAC ACA GTC GCT Thr Gly His Val Ser Arg Leu Ser Pro Leu Ser Val Asp Thr Val Ala 3310	G GTG TCC AGC GTG GTT GTC TAT CCC GTC TAC CTG GCC ATC u Vai Ser Ser Vai Vai Vai Tyr Pro Vai Tyr Leu Aia Iie 3330 3340
CTG GTG GAG AAG GAG GTG CTG Leu Vai Giu Lys Giu Vai Leu 3230	TTC CGG CGC CTG (Phe Arg Arg Leu 3245	CAC ATC TGG C1 His Ile Trp L6	CGC ATC CAG AO Arg Ile GIN AI	GGC GCC AAC GO Gly Ald Ash A 3295	ACG GGG CAT G Thr Gly His V 3310	GTT GGC CTG G Val Gly Leu V 3325

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				. •		FIG. 1
10202	10250	10298	10346	10394	10442	10490
TTT CTC TTC CGG ATG TCC CGG AGC AAG GTG GCT GGG AGC CCG AGC Phe Leu Phe Arg Met Ser Arg Ser Lys Val Ala Gly Ser Pro Ser 3345	A CCT GCC GGG CAG CTG CTG GAC ATC GAC AGC TGC CTG GAC r Pro Aig Gly Gin Gin Val Leu Asp Ile Asp Ser Cys Leu Asp 3360	TCC GTG CTG GAC AGC TCC TTC CTC ACG TTC TCA GGC CTC CAC GCT Ser Val Leu Asp Ser Ser Phe Leu Thr Phe Ser Gly Leu His Ala 3375	T GTT GGA CAG ATG AAG AGT GAC TTG TTT CTG GAT GAT TCT e Vai Gly Gin Met Lys Ser Asp Leu Phe Leu Asp Asp Ser 3395	G GTG TGC TGC CCC TCC GGC GAG GGA ACG CTC AGT TGG CCG u Vai Cys Trp Pro Ser Gly Glu Gly Thr Leu Ser Trp Pro 3410 3420	CTG CTC AGT GAC CCG TCC ATT GTG GGT AGC AAT CTG CGG CAG CTG Leu Leu Ser Asp Pro Ser Iie Val Gly Ser Asn Leu Arg Gln Leu 3435	CGG GGC CAG GCG CGC CAT GGG CTG GGC CCA GAG GAG GAC GGC TTC Arg Gly Gin Aig Gly His Gly Leu Gly Pro Glu Glu Asp Gly Phe 3440
CTT TTT CT Leu Phe Le	CCC ACA CC Pro Thr Pr	TCG TCC GT Ser Ser Vg	GAG GCC TTT GTT Glu Ala Phe Val 3390	AAG AGT CTG G Lys Ser Leu V 3405	GAC CTG CT Asp Leu Le	GCA CGG GG Ala Arg GI

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10538		
TCA GAT	Asp	
TCA	Ser	
GCA	A I d	
TTC TCA GCA	Ser	3465
110	Phe	
ည	Ser	
AAA	Lys	
ည္ဟ	Aid	
CCT	r Pro Aid Lys Ser	3460
1 00	Ser	
TAC	Tyr	
ပ္ပ	Pro Tyr Ser	
AGC	Ser	
	Ald	3455
CTG GCC	Leu Ala	-
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Applicant: Oxana Ibraghimov-Beskrovnaya, et al. Title: Compositions and Methods for Treating Polycystic

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10586 ပ္ပ Pro Ald SCA CAG CAG GTC CTT GCC GAG GGG GTC AGC AGC GIN GIN Val Leu Ala Glu Gly Val Ser Ser 3480 3475 GAC CTG ATC C Asp Leu Ile G GAA

10634 AGC CTG TCC Ser Leu Ser 3500 GAA ACG GAC CTG CTC AGC Glu Thr Asp Leu Leu Ser 3495 ACC CAC ATG (Thr His Met (ATG 3490 GIN ASP GAC CAA Pro Thr 3485 ACC

10682 GAG ACG CTG GCG CTG CAG AGG CTG GGG Glu Thr Leu Ala Leu Gin Arg Leu Gly 3510 GAG AAG ACA GIU Lys Thr 3505 CI y 999 Pro CCT ACT Thr AGC Ser

10730 TGG GAA CAG CCC CAG GCA Trp Glu Gln Pro Gln Ala 3530 , Gly Leu Asn T 3525 CTG AAC ည္ဟ Ser Pro AGC CCA Pro Pro SGG CCA CCC 3520 <u>C</u> | CIG Leu GAG

10778 CTG Leu GCG AGG CTG TCC AGG ACA GGA CTG GTG GAG GGT CTG CGG AAG CGC Ala Arg Leu Ser Arg Thr Gly Leu Val Glu Gly Leu Arg Lys Arg 3535 3545

10826 TCC CTG GCC CAC GGG CTC AGC CTG CTC CTG Ser Leu Ala His Gly Leu Ser Leu Leu Leu 3555 ည္ဟ Ald 100 101 Trp Cys CTG CCG GCC Pro Ald ren

FIG. 1AF

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10874		
ည္ဟ	Pro	3580
	Phe	
AGC	Ser	
909	Ald	
SCT CCG	<u>C</u>	
GTG	D/	3575
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310	0	
GCT	AIG	3570
GTG	ΙDΛ	
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Polycystic Kidney Disease

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10922 TTC CTG Phe Leu AGC GTT GCG TGG CTC CTG TCC AGC AGC GCC Val Ala Trp Leu Leu Ser Ser Ser Ala 3590 3585 Ser AGT GTG <u>|</u> ၁၅၅ <u>GI y</u> ၅၁၁ GTG Val 3565

10970 CTG Trp Giu Pro Leu Lys Vai Leu Leu Giu Ala Leu 3605 GAA GCC CTG AAG GTC TTG CTG SCA GAG 166 166 Phe Leu Gly CTC 66C 3600 Ser TCA

11018 ACC Pro Asp Glu Asp Asp Thr 3625 CCG GAT GAA GAT GAC Arg Leu His 1 3620 CGC CTG CAC TCA CTG GTG GCC AAG Ala Lys Ser Leu Val / 3615 Phe TAC Tyr

11066 AGC GCA CGT GTG CCC CGC Ser Ald Arg Val Pro Arg 3640 GCT GTG ACG CCT GTG AV Ala Val Thr Pro Val S 3635 CCG GCT Pro GTA GAG AGC (Val Glu Ser F 3630 Leu

11114 3660 Ald GAA GAA GCC n S <u>n</u>19 CTG GCC AAG (Leu Ala Lys (3655 CAC GGC TTT GCA CTC TTC His Gly Phe Ala Leu Phe 3650 CCA CCC Pro I Arg CTA CGG Val ,3645

11162 GTG TAC Vai Tyr 3675 CTG CGG AGC CTC CTG G Leu Arg Ser Leu Leu V 3670 AAG GTC AAG AGG CTA CAT GGC ATG Lys Val Lys Arg Leu His Gly Met Arg Leu 3665 ပ္ပ

FIG. 1AG

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ATC CTT TIT CTC CTC GTC ACC CTC GCC ACC TAT GGC GAT GCC TCA 11210 Met Leu Phe Leu Leu Val Thr Leu Leu Ala Ser Tyr Gly Asp Ala Ser 3680 TGC CAT GGC CAC CTC CGT CTC CAA AGC GCC ATC AAG CAG CTG 11258 CAC AGC CGC CAC CTC CGT CTC CAA AGC GCC ATC AAG CAG CAG CTG 11306 His Ser Arg Ala Tyr Arg Leu Gln Ser Ala IIe Lys Gln Glu Leu Synos TGG ATG CCC CAC CTC CTG CAC CGC TTC AGG CAG CTC TGG CCA 11306 His Ser Arg Ala Phe Leu Ala IIe Thr Arg Ser Glu Glu Leu Trp Pro 3710 TGG ATG CCC CAC CTG CTG CTG CTG CTG CAG CAG CAG CAG CAG TTC AGG CAG CTG CAG CAG CTG CAG CAG CTG CTG CAG CTG CAG CTG CTG CAG CTG CTG CAG CTG CTG CAG CTG CTG CTG CTG CTG CTG CTG CTG CTG CT	APR 0 1 2003 &		7	35/59			FIG. 1AH
TG ACC CTG CTG GCC AGC TAT GGG Thr Leu Leu Alg Ser Tyr Gly 3685 AC CGT CTG CAA AGC GCC ATC AAG yr Arg Leu Gin Ser Alg Ile Lys 3700 TG GCC ATC ACG CGG TCT GAG GAG eu Alg Ile Thr Arg Ser Glu Glu Glu S715 TG CTG CCC TAC GTC CAC GGG AAC eu Leu Pro Tyr Val His Gly Asn 3730 CA CGG CTG CGG CAG GTG CGG CTG CA CGG CTG CGG CAT TAC GAC GTT GGC TGG CAG CTG CAT TAC CAC GTT CGG CTG	11210	11258	11306	11354	11402	11450	11498 F
	rg ACC CTG CTG GCC AGC TAT GGG GAT GCC al Thr Leu Leu Ala Ser Tyr Gly Asp Ala 3685	AC CGT CTG CAA AGC GCC ATC AAG CAG GAG yr Arg Leu Gin Ser Aig Iie Lys Gin Giu 3700	AC AGC CGG GCC TTC CTG GCC ATC ACG CGG TCT GAG GAG CTC TGG CCA is Ser Arg Ald Phe Leu Ald Ile Thr Arg Ser Glu Glu Leu Trp Pro 3710 3710	CAC GGG AAC His Gly Asn 3735	STG GGG CCC CCA CGG CTG CGG CAG GTG CGG CTG Leu Gly Pro Pro Arg Leu Arg Gln Val Arg Leu 3745	TC TAC CCA GAC CCT CCC GGC CCC AGG GTC CAC ACG TGC TCG GCC GCA eu Tyr Pro Asp Pro Pro Gly Pro Arg Val His Thr Cys Ser Ala Ala 3760 3765	TTC AGC ACC AGC GAT TAC GAC GTT Phe Ser Thr Ser Asp Tyr Asp Val 3775

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	APR 0 1 2003	648		36/59			1AI
	11546	11594	11642	11690	11738	11786	FIG. 1AI
Kidney Disease Page 36 of 59	AAT GGC TCG GGG ACG TGG GCC TAT TCA GCG CCG GAT CTG CTG GGG GCA ASN GIY Ser GIY Thr Trp AIG TYr Ser AIG Pro ASP Leu Leu GIY AIG 3790	TGG TCC TGG GGC TCC TGT GCC GTG TAT GAC AGC GGG GGC TAC GTG CAG Trp Ser Trp Gly Ser Cys Ala Val Tyr Asp Ser Gly Gly Tyr Val Gln 3805	GAG CTG GGC CTG AGC CTG GAG AGC CGC GAC CGG CTG CGC TTC CTG Glu Leu Gly Leu Ser Leu Glu Glu Ser Arg Asp Arg Leu Arg Phe Leu 3835	CAG CTG CAC AAC TGG CTG GAC AAC AGG AGC CGC GCT GTG TTC CTG GAG GIN Leu His Asn Trp Leu Asp Asn Arg Ser Arg Ala Val Phe Leu Glu 3840	CTC ACG CGC TAC AGC CCG GCC GTG GGG CTG CAC GCC GCC GTC ACG CTG Leu Thr Arg Tyr Ser Pro Ala Val Gly Leu His Ala Ala Val Thr Leu 3855	CGC CTC GAG TTC CCG GCG GCC GGC GCC CTG GCC GCC CTC AGC GTC Arg Leu Giu Phe Pro Aia Aia Giy Arg Aia Leu Aia Aia Leu Ser Vai 3870 3875	CGC CCC TTT GCG CTG CGC CTC AGC GCG GGC CTC TCG CTG CTT CTG Arg Pro Phe Ald Leu Arg Arg Leu Ser Ald Gly Leu Ser Leu Pro Leu 3885

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APR 0 1 2003	04837		37/59		1AJ			
11882	11930	11978	12026	12074	12122	12170 FIG. 1AJ		
CTC ACC TCG GTG TGC CTG CTG TTC GCC GTG CAC TTC GCC GTG GCC Leu Thr Ser Val Cys Leu Leu Leu Phe Ala Val His Phe Ala Val Ala 3905	GAG GCC CGT ACT TGG CAC AGG GAA GGG CGC TGG CGC GTG CTG CGG CTC Glu Ala Arg Thr Trp His Arg Glu Gly Arg Val Arg Val Leu Arg Leu 3920	GGA GCC TGG CGG TGG CTG CTG GCG CTG ACG GCG GCC ACG GCA Gly Ala Trp Ala Arg Trp Leu Leu Val Ala Leu Thr Ala Ala Thr Ala 3935	CTG GTA CGC CTC GCC CAG CTG GGT GCC GCT GAC CGC CAG TGG ACC CGT Leu Vai Arg Leu Aia Gin Leu Giy Aia Aia Asp Arg Gin Trp Thr Arg 3950	TTC GTG CGC CGC CCG CGC TTC ACT AGC TTC GAC CAG GTG GCG Phe Val Arg Gly Arg Pro Arg Arg Phe Thr Ser Phe Asp Gln Val Ala 3965 3970 3970	GAG CTG AGC TCC GCA GCC CGT GGC CTG GCG GCC TCG CTG CTC TTC CTG GIN Leu Ser Ser Ala Ala Arg Gly Leu Ala Ala Ser Leu Leu Phe Leu 3985	CTT TTG GTC AAG GCT GCC CAG CTA CGC TTC GTG CGC CAG TGG TCC Leu Leu Vai Lys Aia Aia Gin Gin Leu Arg Phe Vai Arg Gin Trp Ser 4000		

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TRADEMARK CHI	1048		38/59		·	FIG. 1AK		
12218	12266	12314	12362	12410	12458	12506 FI(
GTC TTT GGC AAG ACA TTA TGC CGA GCT CTG CCA GAG CTC CTG GGG GTC Vai Phe Giy Lys Thr Leu Cys Arg Aia Leu Pro Giu Leu Leu Giy Vai 4015 4015	ACC TTG GGC CTG GTG CTC GGG GTA GCC TAC GCC CAG CTG GCC ATC Thr Leu Gly Leu Val Val Leu Gly Val Ala Tyr Ala Gln Leu Ala Ile 4030	CTG CTC GTG TCT TCC TGT GTG GAC TCC CTC TGG AGC GTG GCC CAG GCC Leu Leu Vai Ser Ser Cys Vai Asp Ser Leu Trp Ser Vai Ala Gin Ala 4045	CTG TTG GTG CTG TGC CCT GGG ACT GGG CTC TCT ACC CTG TGT CCT GCC Leu Leu Vai Leu Cys Pro Gly Thr Gly Leu Ser Thr Leu Cys Pro Ala 4075	GAG TCC TGG CAC CTG TCA CCC CTG CTG TGT GTG GGG CTC TGG GCA CTG Glu Ser Trp His Leu Ser Pro Leu Leu Cys Val Gly Leu Trp Ala Leu 4080	CGG CTG TGG GGC GCC CTA CGG CTG GGG GCT GTT ATT CTC CGC TGG CGC Arg Leu Trp Giy Aig Leu Arg Leu Giy Aig Vai Iie Leu Arg Trp Arg 4105	TAC CAC GCC TTG CGT GGA GAG CTG TAC CGG CCG GCC TGG GAG CCC CAG Tyr His Ala Leu Arg Gly Glu Leu Tyr Arg Pro Ala Trp Glu Pro Gln 4110		

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	12554	12602	12650	12698	12746	12794	12842 FIG. 1AI
Title: Compositions and Methods for Treating Polycystic Kidney Disease Page 39 of 59	S CGC AGG J Arg Arg 4135	GGC CTC AGC AAG GAG TTC CGC CAC AAA GTC CGC TTT GAA GGG Gly Leu Ser Lys Val Lys Glu Phe Arg His Lys Val Arg Phe Glu Gly 4155	ATG GAG CCG CTG CCC TCT CGC TCC TCC AGG GGC TCC AAG GTA TCC CCG Met Giu Pro Leu Pro Ser Arg Ser Ser Arg Gly Ser Lys Val Ser Pro 4160	GAT GTG CCC CCA CCC AGC GCT GGC TCC GAT GCC TCG CAC CCC TCC ACC Asp Val Pro Pro Pro Ser Ala Gly Ser Asp Ala Ser His Pro Ser Thr 4175	TCC TCC AGC CAG CTG GAT GGG CTG AGC GTG AGC CTG GGC CGG CTG GGG Ser Ser Ser Gin Leu Asp Gly Leu Ser Val Ser Leu Gly Arg Leu Gly 4190	ACA AGG TGT GAG CCT GAG CCC TCC CGC CTC CAA GCC GTG TTC GAG GCC Thr Arg Cys Glu Pro Glu Pro Ser Arg Leu Gln Alg Val Phe Glu Alg 4205	CTG CTC ACC CAG TTT GAC CGA CTC AAC CAG GCC ACA GAG GAC GTC TAC Leu Leu Thr Gin Phe Asp Arg Leu Asn Gin Aig Thr Giu Asp Vai Tyr 4235

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Polycystic Kidney Disease

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999	. Arg	
AGC	Ser	_
AGC	Ser	4250
AGG	Arg	ı
၁၅၁	Arg	
၁၅၅	Ser Leu Gin Giy Arg Arg Ser Ser	
CAA	GIn	
CTG	Leu	4245
AGC	Ser	
CAC	His	
CAG CTG CAC	Leu	
CAG	C U	_
CAG	Gun	4240
GAG	n 19	
CTG	Gin Leu Giu Gin	
CAG	GIn	

12938 4260

12986 CCC AGC CGC CTT GCC CGG GCC AGT CGG GGT GTG GAC CTG GCC ACT Pro Ser Arg Leu Ala Arg Ala Ser Arg Gly Val Asp Leu Ala Thr 4270 Leu

13034 4300 GGC CCC AGC AGG ACA CCC CTT CGG GCC AAG AAC AAG GTC CAC CCC AGC GIY Pro Ser Arg Thr Pro Leu Arg Aid Lys Asn Lys Val His Pro Ser 4290

13090 AGC ACT TAGTCCTCCT TCCTGGCGGG GGTGGGCCGT GGAGTCGGAG TGGACACCGC Ser Thr TCAGTATTAC TITCTGCCGC TGTCAAGGCC GAGGGCCAGG CAGAATGGCT GCACGTAGGT 13150

TCCCCAGAGA GCAGGCAGGG GCATCTGTCT GTCTGTGGGC TTCAGCACTT TAAAGAGGCT 13210

GTGTGGCCAA CCAGGACCCA GGGTCCCCTC CCCAGCTCCC TTGGGAAGGA CACAGCAGTA 13270

TTGGACGGTT TCTAGCCTCT GAGATGCTAA TTTATTTCCC CGAGTCCTCA GGTACAGCGG 13330

FIG. 1AM



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14060 13570 CAAGGCCCTC GGAGCTGGCT GTGCCTGCTT CTGTGTACCA CTTCTGTGGG CATGGCCGCT 13990 TCTAGAGCCT CGACACCCCC CCAACCCCCG CACCAAGCAG ACAAAGTCAA TAAAAGAGCT 14050 AGGGIGGTIA GAGGAAAAGA CTCCTCCTGG GGCCTGGCTC CCAGGGTGGA GGAAGGTGAC 13870 TOTOTOTOTO TOTOTOTOCO COCOCCACO COCOAGIGIO CIGIATOCOC CAGOCAGOCT 13930 GGAGGGTTAG CCTGCACCGC CGCCACCCTG CCCCTAAGTT ATTACCTCTC CAGTTCCTAC 13450 CCTCCCCCAA CACCTGCTGC GCTTGGTAGG TGTGGTGGCG TTATGGCAGC CCGGCTGCTG 13630 CTTGGATGCG AGCTTGGCCT TGGGCCGGTG CTGGGGGCAC AGCTGTCTGC CAGGCACTCT 13690 CATCACCCCA GAGGCCTTGT CATCCTCCCT TGCCCCAGGC CAGGTAGCAA GAGAGCAGCG 13750 CCCAGGCCTG CTGGCATCAG GTCTGGGCAA GTAGCAGGAC TAGGCATGTC AGAGGACCCC 13810 GCTGTGCCCG GCCCCACCCC CTGGGCAGAT GTCCCCCACT GCTAAGGCTG CTGGCTTCAG 13390 CGTACTCCCT GCACCGTCTC ACTGTGTGTC TCGTGTCAGT AATTTATATG GTGTTAAAAT GTGTATATTT TTGTATGTCA CTATTTTCAC TAGGGCTGAG GGGCCTGCGC CCAGAGCTGG GTCTGACTGC FIG. 1AN



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specific		PKD1	FP-L1 Res. 2621-2710	FP-L2 Res. 2734-3094	FP-L3 Res. 3116-3300	FP-L4 Res. 3364-3578	FP-L5 Res. 3623-3688	FP-L6 Res. 3710-3914	FP-L7 Res. 3931-4046	MAL-REJ Res. 2166-2599	MAL-BD3 Res. 4097-4302] FP-46-2 Res. 4148-4219	FP-46-1c Res. 4220-4302	FP-LRR Res. 27-360
Polyclonal antibodies to domain spe polycystin fusion proteins	S LRR Ig-like REJ	1 3 4 5 7												
			CST	GST	GST	GST	GST	GST	GST	MBP	MBP	CST	GST	GST

 ${
m FIG}^{\sf v}~2$



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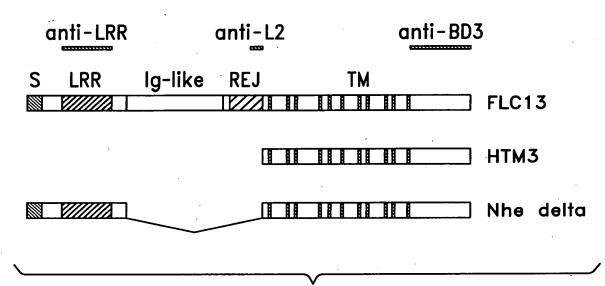
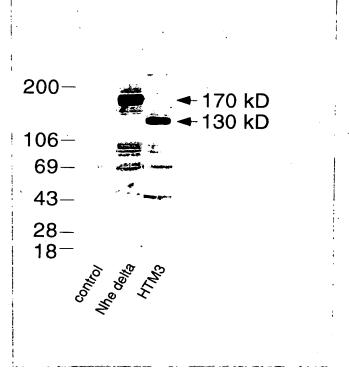


FIG. 3A

FIG. 3B





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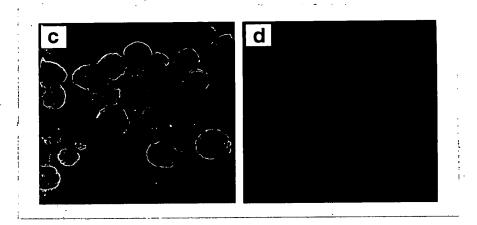
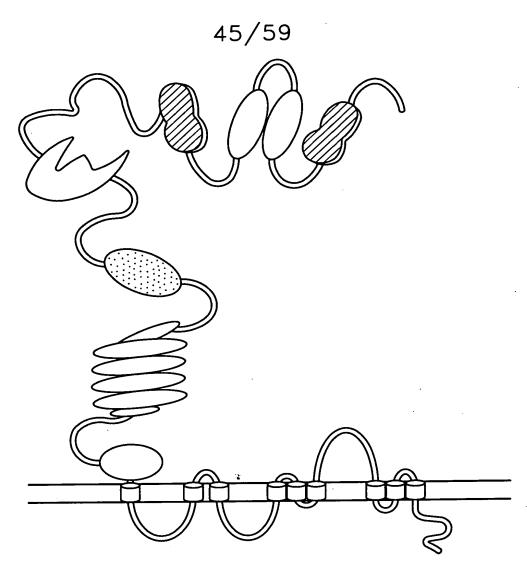


FIG. 3D **3**C FIG.



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N-amino flanking region C-carboxy flanking region



LRR - leucine - rich repeats



→ Ig-like domains



<equation-block> C-type lectin domain



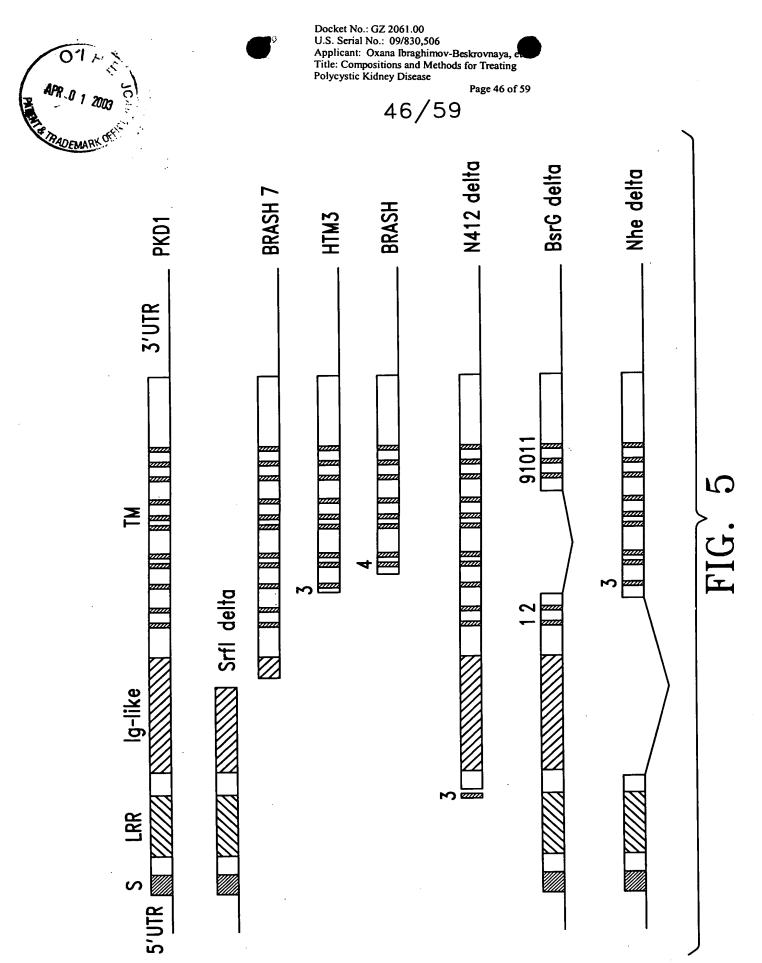
REJ - domain with homology to the receptor for egg jelly



LDL - like domain



TM-putative transmembrane region





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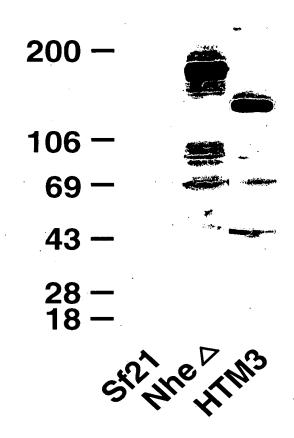


FIG. 6



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S TM LRR S 3'UTR Nhe delta 5'UTR

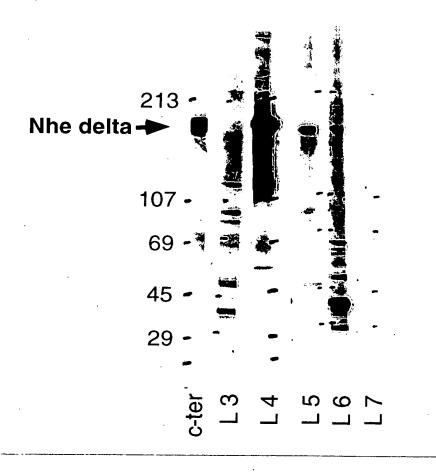


FIG. 7



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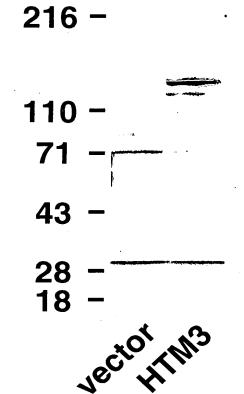


FIG. 8



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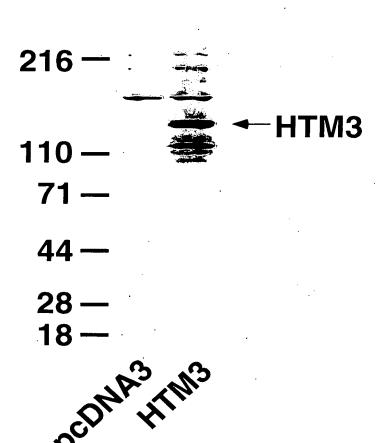


FIG. 9

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→ in vitro translated kidney 69



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Na∕K[†] ATPase PKD1 PKD1 Applicant: Oxana Ibraghimov-Beskrovnaya, et al. litle: Compositions and Methods for Treating Page 52 of 59 200 Polycystic Kidney Disease 106 69 43 28 18 kidney brain brain

FIG. 10B



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kidney

brain

28-

FIG. 10C

14



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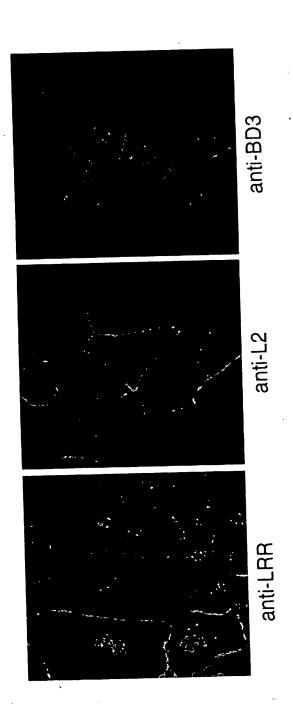
199 106 69 43 28 -

FIG. 10D



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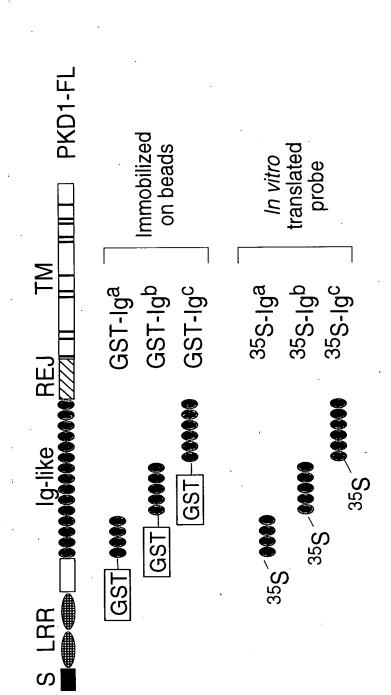




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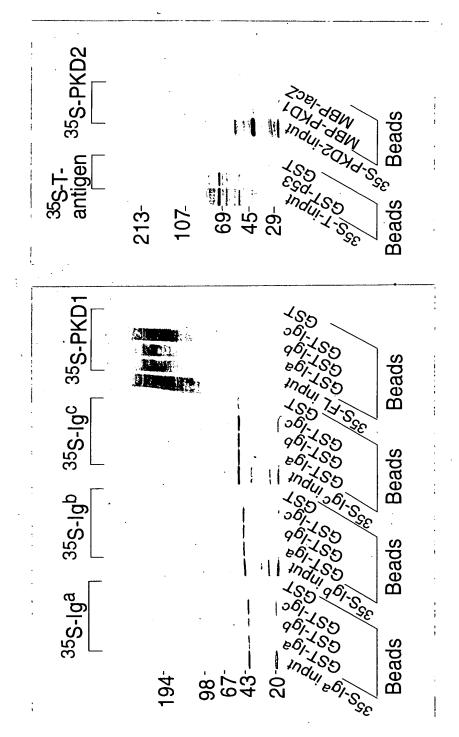


FIG. 12C

FIG. 12B



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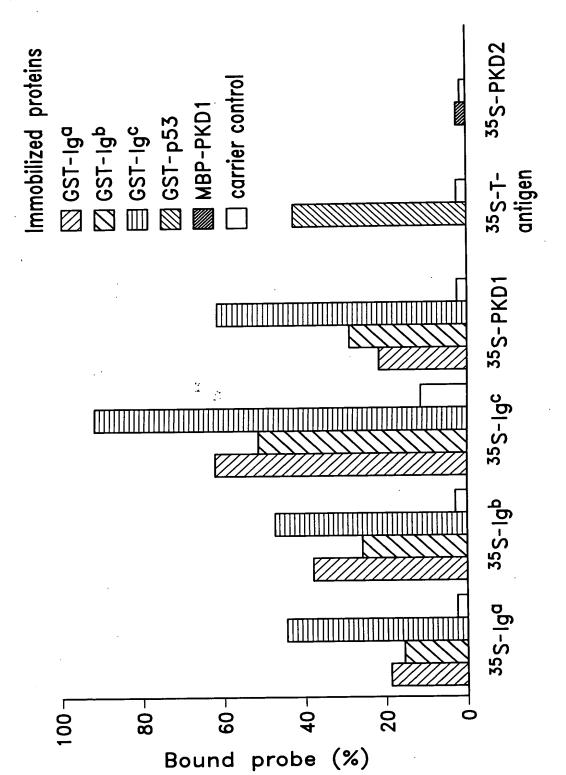


FIG. 13



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